

FIGURE 1

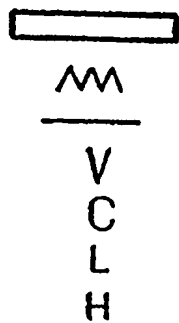
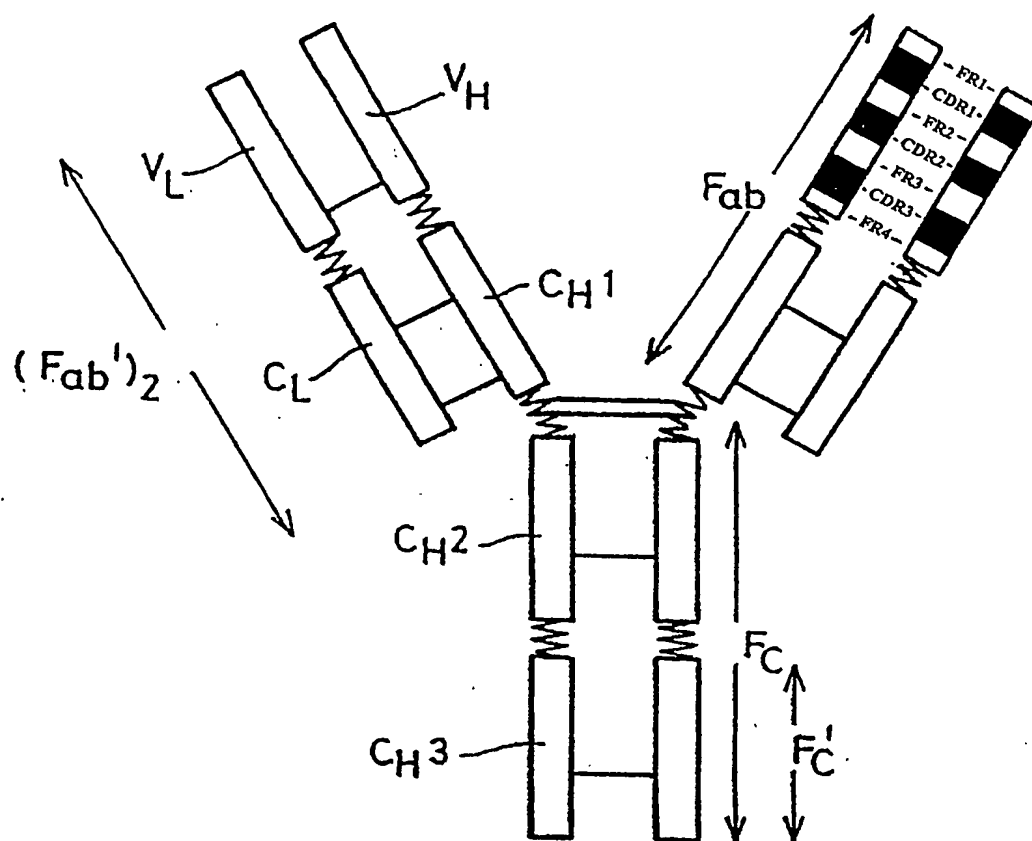


FIGURE 2**A.****SEQ ID NO:59 - AME 33 light chain variable region amino acid sequence**

EIVLTQSPGTL~~SL~~SPGERATL~~SCR~~ASSSV~~PI~~HWYQQKPGQAP~~RL~~LIYATSALASGI
PDRFSGSGSGTDFTLTISRLEPEDFAVYYCQQWLSNPPTFGQGTKLEIK

B.**SEQ ID NO:60 - AME 33 light chain variable region nucleic acid sequence**

GAAATTGTGTTGACGCAGTCTCCAGGCACCCTGTCTTTGTCTCCAGGGGAAAG
AGCCACCCTCTCCTGCAGGGCCAGCTCAAGTGTACCGTACATCCACTGGTAC
CAGCAGAAACCTGGCCAGGCTCCCAGGCTCCTCATCTATGCCACATCCGCTCT
GGCTTCTGGCATCCCAGACAGGTTCAAGTGGCAGTGGGTCTGGGACAGACTTC
ACTCTCACCATCAGCAGACTGGAGCCTGAAGATTTTGCAGTGTATTACTGTCA
GCAGTGGCTGAGTAACCCACCCACTTTTGGCCAGGGGACCAAGCTGGAGATC
AAA

FIGURE 3**A.****SEQ ID NO:61 - AME 33 heavy chain variable region amino acid sequence**

EVQLVQSGAEVKKPGESLKISCKGSGRTFTSYNMHWVRQMPGKGLEWMGAIYP
LTGDTSYNQKSKLQVTISADKSISTAYLQWSSLKASDTAMY YCARSTYVGGDW
QFDVWGKGTTVTVSS

B.**SEQ ID NO:62 - AME 33 heavy chain variable region nucleic acid sequence**

GAGGTGCAGCTGGTGCAGTCTGGAGCAGAGGTGAAAAAGCCCGGGGAGTCT
CTGAAGATCTCCTGTAAGGGTTCTGGCCGTACATTTACCAGTTACAATATGCA
CTGGGTGCGCCAGATGCCCGGGAAAGGCCTGGAGTGGATGGGGGCTATTTAT
CCCTTGACGGGTGATACTTCCTACAATCAGAAGTCGAAACTCCAGGTCACCA
TCTCAGCCGACAAGTCCATCAGCACCGCCTACCTGCAGTGGAGCAGCCTGAA
GGCCTCGGACACCGCCATGTATTACTGTGCGAGATCGACTTACGTGGGCGGT
GACTGGCAGTTCGATGTCTGGGGCAAGGGGACCACGGTCACCGTCTCCTCA

FIGURE 4**A. Amino acid sequence of a human light chain framework region VkIII (A27) (DPK22) with interspersed CDR sequences labeled**

(FRL1 - SEQ ID NO:71) CDRL1 (FRL2 - SEQ ID NO:72) CDRL2
 EIVLTQSPGTLSPGERATLSCXXXXXXXXXXXXWYQQKPGQAPRLLIYXXXXXXXXXX
 (FRL3 - SEQ ID NO:73) CDRL3 (FRL4 - SEQ ID NO:74)
 GIPDRFSGSGSGTDFTLTISRLEPEDFAVYYCXXXXXXXXXXXXFGQGTKLEIK

B. Nucleic acid sequence of a human light chain framework region VkIII (A27) (DPK22) with interspersed CDR sequences labeled

(FRL1 - SEQ ID NO:75)
 GAAATTGTGTTGACGCAGTCTCCAGGCACCCTGTCTTTGTCTCCAGGGGAAAGAG
 CDRL1
 CCACCCTCTCCTGCXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXTGGTACCAG
 (FRL2 - SEQ ID NO:76) CDRL2
 CAGAAACCTGGCCAGGCTCCCAGGCTCCTCATCTATXXXXXXXXXXXXXXXXXXXX
XXXGGCATCCCAGACAGGTTCAGTGGCAGTGGGTCTGGGACAGACTTCACTCTC
 (FRL3 - SEQ ID NO:77)
 ACCATCAGCAGACTGGAGCCTGAAGATTTTGCAGTGTATTACTGTXXXXXXXXXX
 CDRL3 (FRL4 - SEQ ID NO:78)
XXXXXXXXXXXXXXXXXXXXTTTGGCCAGGGGACCAAGCTGGAGATCAAA

FIGURE 5**A. Amino acid sequence of a human heavy chain framework region VH5-51
(DP-73) with interspersed CDR sequences labeled**

(FRH1 - SEQ ID NO:79) CDRH1 (FRH2 - SEQ ID NO:80)
 EVQLVQSGAEVKKPGESLKISCKGSXXXXXXXXXXWVRQMPGKGLEWMG

CDRH2 (FRH3 - SEQ ID NO:81)
 XXXXXXXXXXXXXXXXXXXXQVTISADKSISTAYLQWSSLKASDTAMYYCAR

CDRH3 (FRH4 - SEQ ID NO:82)
 XXXXXXXXXXXXXXXXWGKGTTVTVSS

**B. Nucleic acid sequence of a human heavy chain framework region VH5-51
(DP-73) with interspersed CDR sequences labeled**

(FRH1 - SEQ ID NO:83)
 GAGGTGCAGCTGGTGCAGTCTGGAGCAGAGGTGAAAAAGCCCGGGGAGTCTCTG

CDRH1
 AAGATCTCCTGTAAGGGTTCTXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXX

(FRH2 - SEQ ID NO:84)
 TGGGTGCGCCAGATGCCCGGGAAAGGCCTGGAGTGGATGGGG

CDRH2
 XXX

CAGGTCACCATCTCAGCCGACAAGTCCATCAGCACCGCCTACCTGCAGTGGAGC

(FRH3 - SEQ ID NO:85)
 AGCCTGAAGGCCTCGGACACCGCCATGTATTACTGTGCGAGAXXXXXXXXXXX

CDRH3 (FRH4 - SEQ ID NO:86)
 XXXXXXXXXXXXXXXXXXXXXXXXXXXXXTGGGGCAAGGGGACCACGGTCACCGTCT

CCTCA

FIGURE 6**A.****SEQ ID NO:63 - AME 5 light chain variable region amino acid sequence**

DIQMTQSPSSLSASVGDRVTITCRASSSVHYIHWYQQKPGKVPKLLIYATSGLAS
GVPSRFSGSGSGTDFTLTISLQPEDVATYYCQWTWTFNPPTFGGGTKVEIK

B.**SEQ ID NO:64 - AME 5 light chain variable region nucleic acid sequence**

GACATCCAGATGACCCAGTCTCCATCCTCCCTGTCTGCATCTGTAGGA
GACAGAGTCACCATCACTTGCAGGGCCAGCTCAAGTGTACATTACATC
CACTGGTACCAGCAGAAACCAGGGAAAGTTCCTAAGCTCTTGATCTAT
GCCACATCCGGCCTGGCTTCTGGGGTCCCATCTCGGTTCAAGTGGCAGT
GGATCTGGGACAGATTTCACTCTCACCATCAGCAGCCTGCAGCCTGAA
GATGTTGCCACTTATTACTGCCAGACTTGGACTTTTAACCCTCCCACG
TTCGGCGGAGGGACCAAGGTGGAGATCAAA

FIGURE 7**A.****SEQ ID NO:65 - AME 5 heavy chain variable region amino acid sequence**

QVQLVQSGAEVKKPGASVKVSCKASGYTFTSYNMHWVRQAPGQGLEWMGAIY
PGNGDTSYNQKFKWRVTMTRDTSTSTVYMELSSLRSEDTAVYYCARSTYYGGD
WQFDEWGKGTTVTVSS

B.**SEQ ID NO:66 - AME 5 heavy chain variable region nucleic acid sequence**

CAGGTGCAGCTGGTGCAGTCTGGTGCTGAAGTGAAGAAGCCTGGGGCC
TCAGTGAAGGTGTCCTGCAAGGCATCTGGATACACCTTCACCAGCTAC
AATATGCACTGGGTGCGACAGGCCCTGGACAAGGGCTTGAGTGGATG
GGAGCCATCTATCCTGGAAATGGTGATACAAGCTACAATCAGAAGTTT
AAATGGAGAGTCACCATGACCAGGGACACGTCCACGAGCACAGTCTAC
ATGGAGCTGAGCAGCCTGAGATCTGAGGACACGGCCGTGTATTACTGT
GCGAGATCGACTTATTACGGCGGTGACTGGCAGTTCGACGAGTGGGGC
AAAGGGACCACGGTCACCGTCTCCTCA

FIGURE 8**A. Amino acid sequence of a human light chain framework region VkI (DPK4) (A20) with interspersed CDR sequences labeled**

(FRL1 - SEQ ID NO:87) (CDRL1) (FRL2 - SEQ ID NO:88) (CDRL2)
 DIQMTQSPSSLSASVGDRVTITCXXXXXXXXXXWYQQKPGKVPKLLIYXXXXXXXX
 (FRL3 - SEQ ID NO:89) (CDRL3) (FRL4 - SEQ ID NO:90)
 GVPSRFSGSGSGTDFTLTISSLQPEDVATYYCXXXXXXXXXXFGGGTKVEIK

B. Nucleic acid sequence of a human light chain framework region VkI (DPK4) (A20) with interspersed CDR sequences labeled

(FRL1 - SEQ ID NO:91)
 GACATCCAGATGACCCAGTCTCCATCCTCCCTGTCTGCATCTGTAGGAGACAGAG
 (CDRL1)
 TCACCATCACTTGCXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXTTGGTACCAG
 (FRL2 - SEQ ID NO:92) (CDRL2)
 CAGAAACCAGGGGAAAGTTCCTAAGCTCTTGATCTATXXXXXXXXXXXXXXXXXXXX
 (CDRL2 cont.) (FRL3 - SEQ ID NO:93)
 XXXXGGGGTCCCATCTCGGTTCAAGTGGCAGTGGATCTGGGACAGATTTCACTCT
 CACCATCAGCAGCCTGCAGCCTGAAGATGTTGCCACTTATTACTGCXXXXXXXX
 (CDRL3) (FRL4 - SEQ ID NO:94)
 XXXXXXXXXXXXXXXXXXXXXTTCGGCGGAGGGACCAAGGTGGAGA

TCAAA

FIGURE 9**A. Amino acid sequence of a human heavy chain framework region VHI DP7/21-2 with interspersed CDR sequences labeled**

(FRH1 - SEQ ID NO:95) (CDRH1) (FRH2 - SEQ ID NO:96)
 QVQLVQSGAEVKKPGASVKVSCKASXXXXXXXXXXXXWVRQAPGQGLEWMG
 (CDRH2) (FRH3 - SEQ ID NO:97)
 XXXXXXXXXXXXXXXXXXXXRVMTTRDTSTSTVYMELSSLRSEDTAVYYCAR
 (CDRH3) (FRH4 - SEQ ID NO:98)
 XXXXXXXXXXXXXXXXWGKGTTVTVSS

B. Nucleic acid sequence of a human heavy chain framework region VHI DP7/21-2 with interspersed CDR sequences labeled

(FRH1 - SEQ ID NO:99)
 CAGGTGCAGCTGGTGCAGTCTGGTGCTGAAGTGAAGAAGCCTGGGGCCTCAGTG
 (CDRH1)
 AAGGTGTCCTGCAAGGCATCTXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXXX
 (FRH2 - SEQ ID NO:100) (CDRH2)
 GGTGCGACAGGCCCTGGACAAGGGCTTGAGTGGATGGGAXXXXXXXXXXXXXX
 (CDRH2 cont.)
 XXAGAGTCACCATGAC
 (FRH3 - SEQ ID NO:101)
 CAGGGACACGTCCACGAGCACAGTCTACATGGAGCTGAGCAGCCTGAGATCTGA
 (CDRH3)
 GGACACGGCCGTGTATTACTGTGCGAGAXXXXXXXXXXXXXXXXXXXXXXXXXX
 (CDRH3) (FRH4 - SEQ ID NO:102)
 XXXXXXXXXXXX
 TGGGGCAAAGGGACCACGGTCACCGTCTCCTCA

FIGURE 10**A.****SEQ ID NO:67 - AME 33 complete light chain amino acid sequence**

EIVLTQSPGTL~~SLSPGERATL~~SCRASSSVPYHWYQQKPGQAPRLLIYATS~~ALASGIPDR~~
FSGSGSGTDFTLTISRLEPEDFAVYYCQQWLSNPPTFGQGTKLEIK~~RTVAAPS~~VFIFPPS
DEOLKSGTASVVCLLNNFYPREAKVQWKVDNALQSGNSQESVTEQDSKDSTYSLSST
LTLSKADYEKHKVYACEVTHQGLSSPVTKSFNRGEC

- Constant Region is underlined

B.**SEQ ID NO:68 - AME 33 complete light chain nucleic acid sequence**

GAAATTGTGTTGACGCAGTCTCCAGGCACCCTGTCTTTGTCTCCAGGGGAAAGAG
CCACCCTCTCCTGCAGGGCCAGCTCAAGTGTACCGTACATCCACTGGTACCAGCA
GAAACCTGGCCAGGCTCCCAGGCTCCTCATCTATGCCACATCCGCTCTGGCTTCTG
GCATCCCAGACAGGTTCAAGTGGCAGTGGGTCTGGGACAGACTTCACTCTCACCAT
CAGCAGACTGGAGCCTGAAGATTTTGCAGTGTATTACTGTCAGCAGTGGCTGAGT
AACCCACCCACTTTTGGCCAGGGGACCAAGCTGGAGATCAAACGAACTGTGGCTG
CACCATCTGTCTTCATCTTCCC GCCATCTGATGAGCAGTTGAAATCTGGAAGTACC
TCTGTTGTGTGCCTGCTGAATAACTTCTATCCCAGAGAGGCCAAAGTACAGTGGA
AGGTGGATAACGCCCTCCAATCGGGTAACTCCCAGGAGAGTGTACAGAGCAGGA
CAGCAAGGACAGCACCTACAGCCTCAGCAGCACCCCTGACGCTGAGCAAAGCAGA
CTACGAGAAACACAAAGTCTACGCCTGCGAAGTCACCCATCAGGGCCTGAGCTCG
CCCGTCACAAAGAGCTTCAACAGGGGAGAGTGTTAG

FIGURE 11**A. SEQ ID NO:69 - AME 33 complete heavy chain amino acid sequence**

EVQLVQSGAEVKKPGESLKISCKGSGRTFTSYNMHWVRQMPGKGLEWMGAIYPLTG
 DTSYNQKSKLQVTISADKSISTAYLQWSSLKASDTAMYYCARSTYVGGDWQFDVWG
 KGTTVTVSS**ASTKGPSVFPLAPSSKSTSGGTAALGCLVKDYFPEPVTVSWNSGALTSG**
VHTFPAVLOSSGLYSLSVVTVPSSSLGTQTYICNVNHKPSNTKVDKKVEPKSCDKTH
TCPPCPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYVDGV
EVHNAKTKPREEQYNSTYRVVSVLTVLHODWLNQKEYKCKVSNKALPAPIEKTISKA
KGOPREPOVYTLPPSRDELTKNOVSLTCLVKGFYPSDIAVEWESNGOPENNYKTTTPV
LDSDGSFFLYSKLTVDKSRWQOGNVFSCSVMEALHNHYTQKSLSLSPGK

- Constant Region is underlined; the "D" at position 280 and "K" at position 290 are in bold

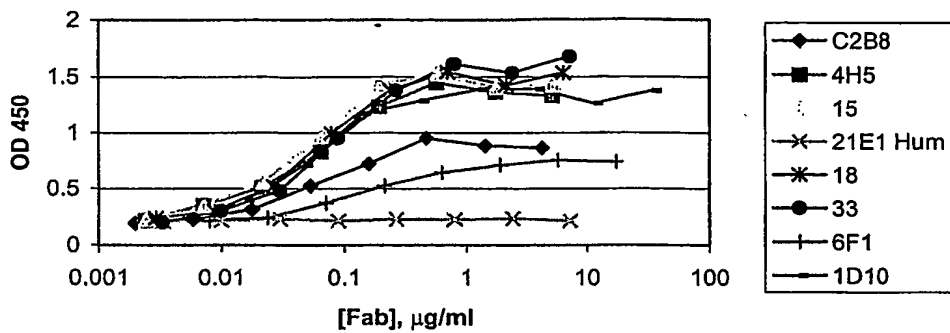
B. SEQ ID NO:70 - AME 33 complete heavy chain nucleic acid sequence

GAGGTGCAGCTGGTGCAGTCTGGAGCAGAGGTGAAAAAGCCCGGGGAGTCTCTG
 AAGATCTCCTGTAAGGGTTCTGGCCGTACATTTACCAGTTACAATATGCACTGGGT
 GCGCCAGATGCCCGGGAAGGCCTGGAGTGGATGGGGGCTATTTATCCCTTGACG
 GGTGATACTTCTACAATCAGAAAGTCGAAACTCCAGGTCACCATCTCAGCCGACA
 AGTCCATCAGCACCGCCTACCTGCAGTGGAGCAGCCTGAAGGCCTCGGACACCGC
 CATGTATTACTGTGCGAGATCGACTTACGTGGGCGGTGACTGGCAGTTCGATGTCT
 GGGGCAAGGGGACCACGGTCACCGTCTCCTCAGCCTCCACCAAGGGCCCATCGGT
 CTTCCCCCTGGCACCCCTCCTCCAAGAGCACCTCTGGGGGCACAGCGGCCCTGGGCT
 GCCTGGTCAAGGACTACTTCCCCGAACCGGTGACGGTGTCTGTGGAAGTCAAGGCGC
 CCTGACCAGCGGCGTGCACACCTTCCCGGTGTCTACAGTCCTCAGGACTCTACT
 CCCTCAGCAGCGTGGTGACCGTGCCCTCCAGCAGCTTGGGCACCCAGACCTACAT
 CTGCAACGTGAATCACAAGCCCAGCAACACCAAGGTGGACAAGAAGGTTGAGCC
 CAAATCTTGTGACAAACTCACACATGCCACCGTGCCAGCACCTGAACTCCTG
 GGGGACCGTCACTCTTCTTCCCCCAAAACCCAAGGACACCCTCATGATCTC
 CCGGACCCCTGAGGTACATGCGTGGTGGTGGACGTGAGCCACGAAGACCCTGAG
 GTCAAGTTCAACTGGTACGTGGACGGCGTGGAGGTGCATAATGCCAAGACAAAG
 CCGCGGGAGGAGCAGTACAACAGCACGTACCGTGTGGTCAGCGTCCTACCGTCC
 TGCACCAGGACTGGCTGAATGGCAAGGAGTACAAGTGCAAGGTCTCCAACAAAG
 CCCTCCCAGCCCCCATCGAGAAAACCATCTCCAAGGCCAAAGGGCAGCCCCGAGA
 ACCACAGGTGTACACCCTGCCCCCATCCCGGACGAGCTGACCAAGAACCAGGTC
 AGCCTGACCTGCCTGGTCAAAGGCTTCTATCCCAGCGACATCGCCGTGGAGTGGG
 AGAGCAATGGGCAGCCGGAGAACAACTACAAGACCACGCCTCCCGTGCTGGACTC
 CGACGGCTCCTTCTTCTCTATAGCAAGCTCACCGTGGACAAGAGCAGGTGGCAG
 CAGGGGAACGTCTTCTCATGCTCCGTGATGCATGAGGCTCTGCACAACCACTACA
 CGCAGAAGAGCCTCTCCCTGTCTCCGGGTAAATGA

FIGURE 12

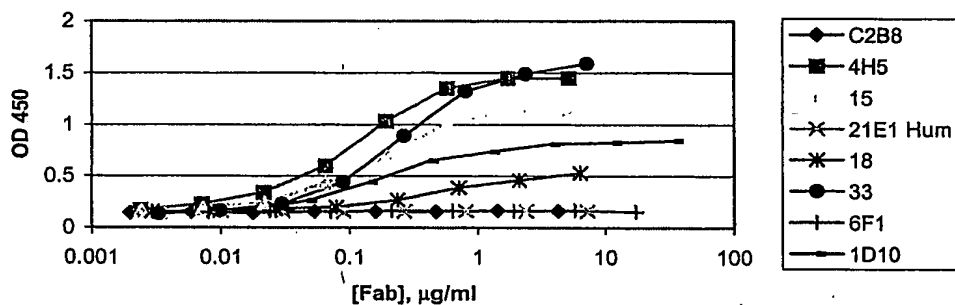
A.

Fab, Binding



B.

Fab, Off rate



C.

Fab, On rate

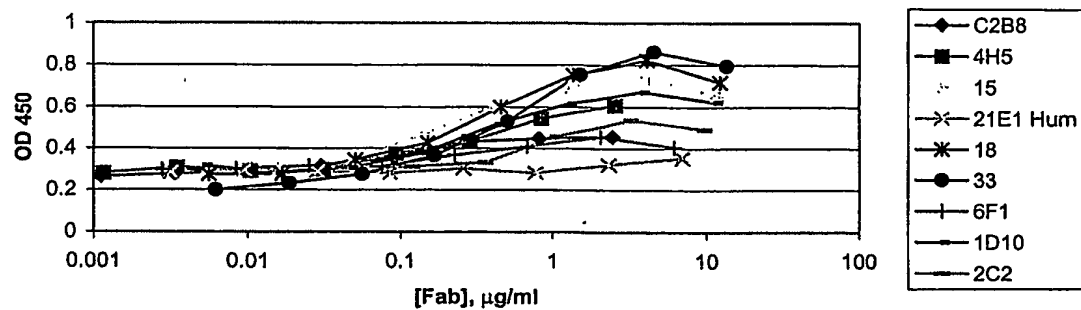
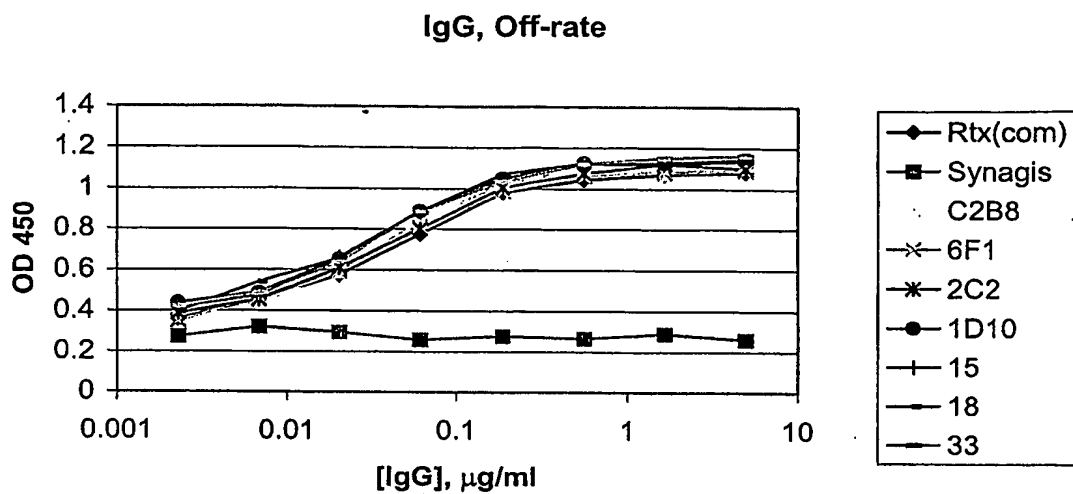


FIGURE 13

A.



B.

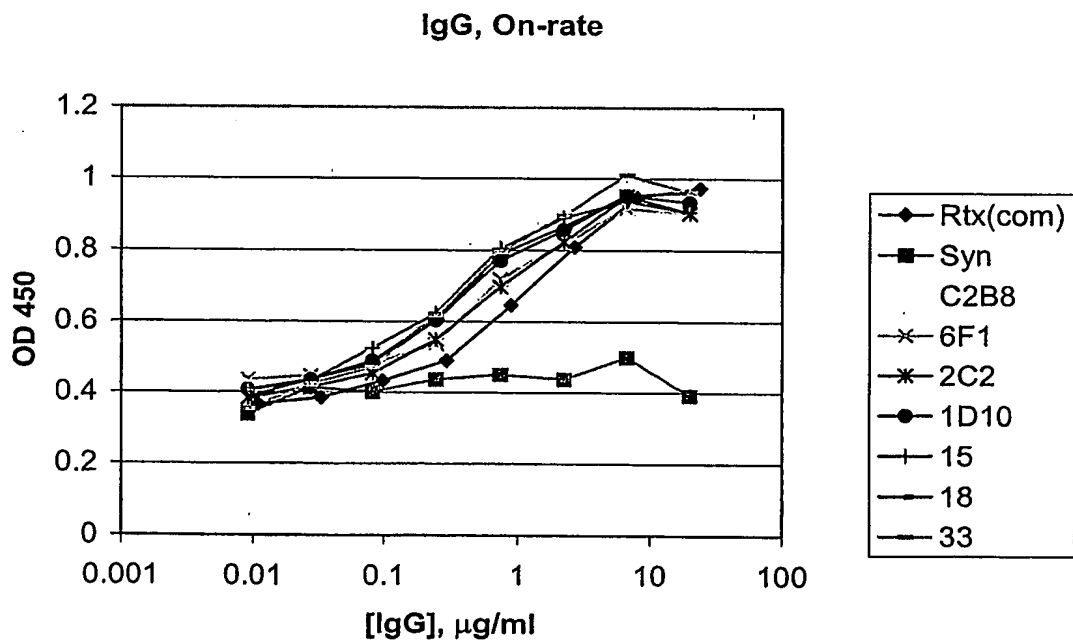
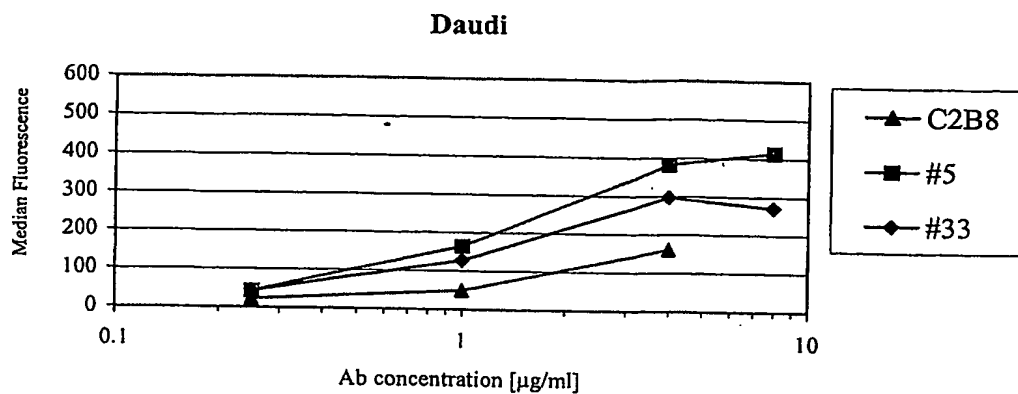
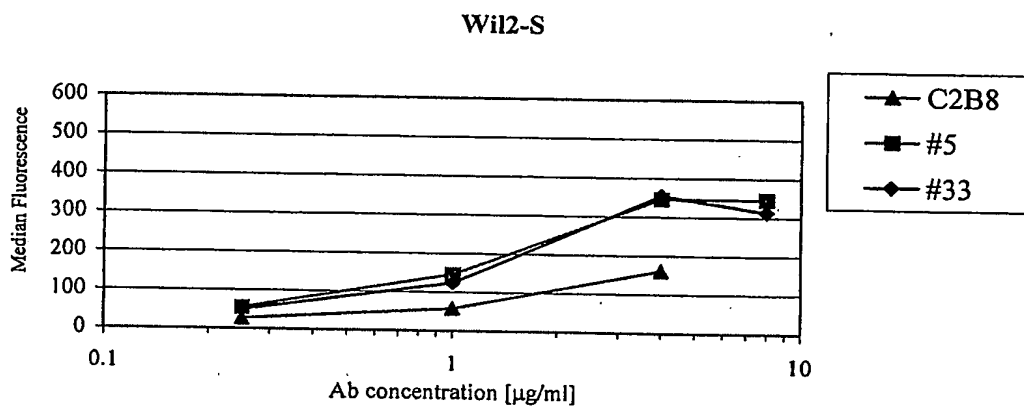


FIGURE 14

A.



B.



C.

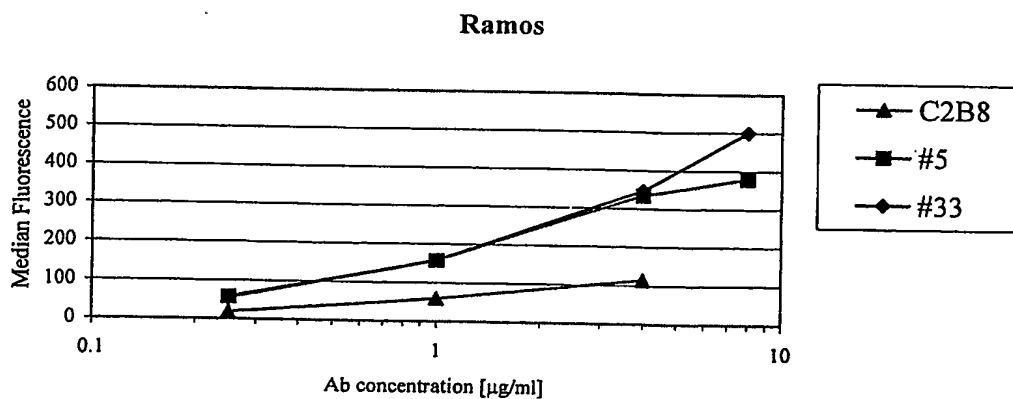


FIGURE 15

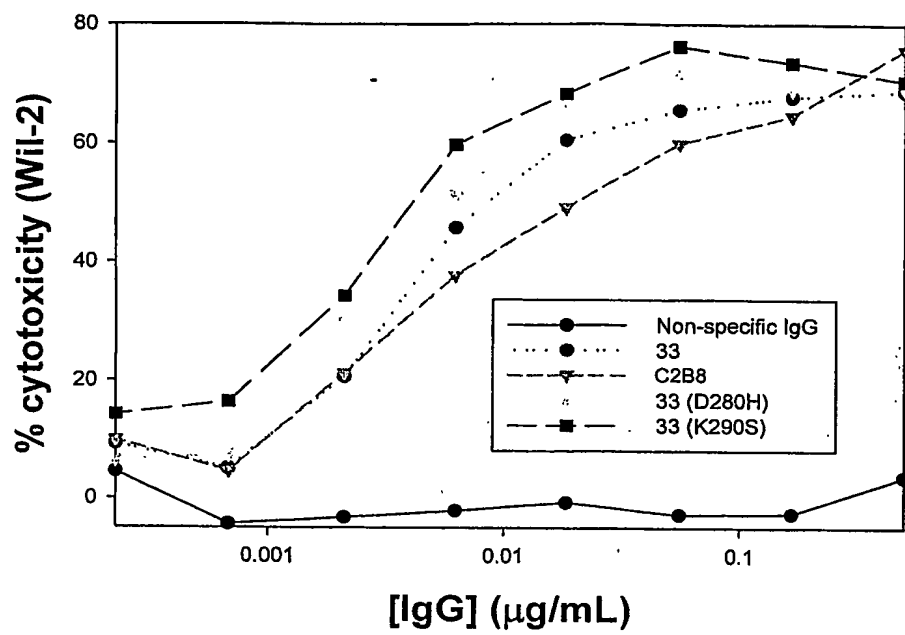
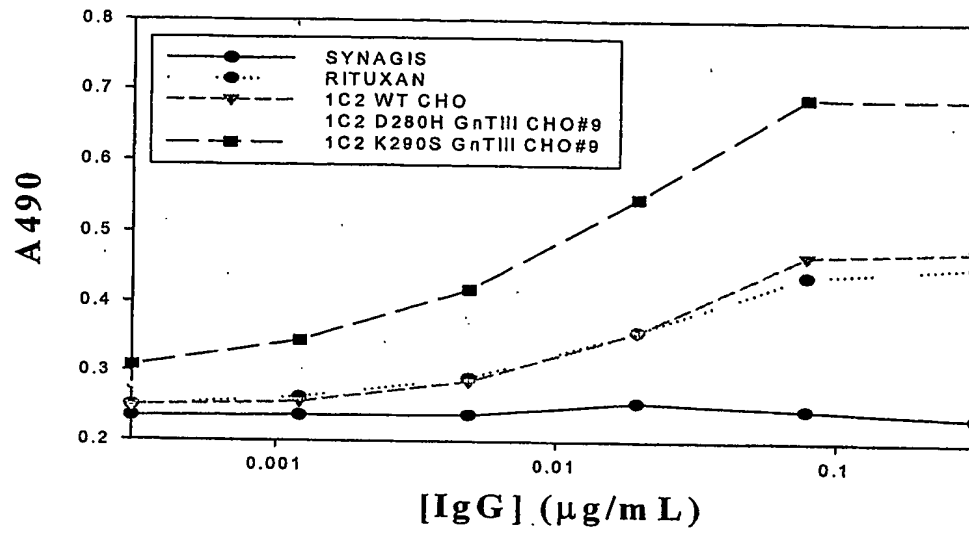


FIGURE 16



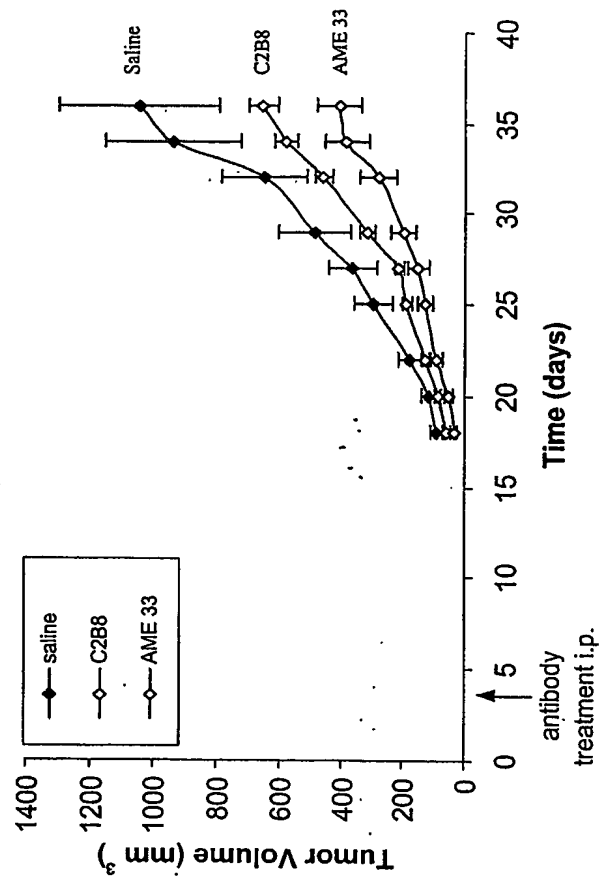


FIGURE 17

17/17